

Fig. 3

SEQ ID NO: 2

MDTSRLGVLL SLPVILLQLAT GGSSPRSGVL LRGCPTHCHC EPDGRMILLRV
DCSDLGLSEL PSNLNSVFTSY LDLSMNNISQ LLPNPLPSLR FLEELRLAGN
5 ALTYIPKGAF TGLYSLKVLM LQNNQLRHVP TEALQNLRSI QSLRLDANHI
SYVPPSCFSG LHSLRHLWLD DNALTEIPVQ AFRSLSALQA MTLALNKIH
IPDYAFGNLS SLVVLHLHNN RIHSLGKKCF DGLHSLETLD LNYNNLDEFP
TAIRTLSNLK ELGFHSNNIR SIPEKAFFVGN PSLITIHFYD NPIQFVGRSA
FQHLPELRTL TLNGASQITE FPDLTGTANL ESLTLTGAQI SSLPQTVCNQ
10 LPNLQVLDLS YNLLEDLPSF SVCQKLQKID LRHNEIYEIK VDTFQQLLSL
RSLNLAWNKI AIIHPNAFST LPSLIKLDLS SNLLSSEPI GLHGLTHLKL
TGNHALQSLI SSENFPELKV IEMPTYAYQCC AFGVCENAYK ISNQWNKGDN
SSMDDLHKKD AGMFQAQDER DLEDFLLDDE EDLKALHSVQ CSPSPGPFKP
CEHLLDGWLI RIGVWTIAVL ALTCNALVTS TVFRSPLYIS PIKLLIGVIA
15 AVNMLTGVSS AVLGVDAFT FGSFARHGAW WENGVGCHVI GFLSIFASES
SVFLLTAAAL ERGFSVKYSA KFETKAPFSS LKVIILLCAL LALTMAAVPL
LGGSKYGASP LCLPLPFGEPE STMGYMVALI LLNSLCFLMM TIAYTKLYCN
LDKGDLHENIW DCSMVKHIAL LLFTNCILNC PVAFLSFSSL INLTFISPEV
IKFILLVVVP LPACLNPLLY ILFNPHFKED LVSLRKQTYV WTRSKHPSLM
20 SINSDDVEKQ SCDSTQALVT FTSSSITYDL PPSSVPSPAY PVTESCHLSS
VAFVPCL

Fig. 4 (1/2)

	LGR5 SEQ ID NO: 2	MDTSRLGVLLSLPVLLQLATGGSSPRSGVLLRGCPTHCHCEPDGRMLLRVDCSDLGSELPSNLSVFTSYLDLSMNNISQ
	LGR6 SEQ ID NO: 3	MRLBGEGRSARAGQNLSRAGSARRGAPR-----DLSMNNLTE
5	LGR4 SEQ ID NO: 4	MPGPLGLLCFLALGLLGSAGPSGAAPLCAAPCSCDGDRR----VDCSGKGLTAVPEGLSAFTQALDISMNNITQ
	Consensus/80%b.L.h.Gpts.tG....sss.pps.sc.....DLSMNN1*p
	LGR5	LLPNPLPSLRFLEELRLAGNALTYPKGAFGLYSLKVIMLQNNQLRHVPTEALQNLRSLQSLRLDANHISYVPPSCFSG
	LGR6	LQPGLFHHLRFLEELRLSGNHLISHIPGQAFTSGLYSLKILMLONNQLGGIPIAEALWELPSLQSL-----D
10	LGR4	LPEDAFKNFPFLLEELQLAGNDLSEIHPKALSGLEKELKVLTQNNQLKTVPSERIRGLSALQSLRLDANHITSVPEDSFEG
	Consensus/80%	L..s.b.pb.FLEELpLtGN.L*a...Ab*GLbpLK1LhLQNNQL.h1PsEAlb.L.tLQSL.....D
	LGR5	LHSLRHLWLDDNALTEIPVQAFRSLSALQMTIALNKIHIIHPDYAFGNLSSLVVLHLHNRRHSLGKKCFDGLHSLETLD
	LGR6	-----D
15	LGR4	LVQLRHLWLDDNSLTVIPVHPLSNLPTLQALTLALKISSIPDEAFTNLSSLVVLHLHNKRSLSQHCFDGLDNLETLD
	Consensus/80%D
	LGR5	LNYYNLDEFPTAIRTLSNLKELGFHHSNNIRSIPEKAFTGNPSLITHFYDNPIQFVGRSAFQHLPELRTLTLNGASQITE
	LGR6	LNYNKLQEFPVAIRTLGRQLQE LGFHNNNNIKAIPEKAFTGMNPPLLQTIHFYDNPIQFVGRSAFQYLPKLHTLSLNGAMDIQE
20	LGR4	LNYYNLGEFPQAIKALPSLKELGFHHSNSISVIPDGAFTGNPLRRTIHYDNPLSFVGNSAFHNLSDLHSLVIRGASMVQQ
	Consensus/80%	LNYNpl.EFP.AI+sLspLpELGFHeNsIpSIP-.AF.GNP.LbTTIbYDNPlpFVGpSAFp.LscL+*Ls1pGA..lpp
	LGR5	FPDLTGTALESLLTGAQISSLPQTVCNQLPNLQVLDLSYNLLEDLPSFSVCQKLQK1DLRNHEIYEIKVDTFQQLLSL
	LGR6	FPDLKGTTSLBILTLTRAGIRLLPSGMCCQQLPRLRVLELSHQNQIEELPSLHRCQKLEEIGLQHNRIWEIGADTFSQLSSL
25	LGR4	FPNLTGTVHLESLLTGTKISSIPNNLCQEOKMLRTLDSYNIRDLPBFNGCHALEEISLQRNQIYQIKEGTFOGLISL
	Consensus/80%	FPsLpGtspLE.LTLT.s.Ip..1PpshCpb..LpsL-LSaN,lc-LPsbp.Cp.LpcIsLp+NpIapI..sTfp.L.SL
	LGR5	RSLNLAWNKIAIIHPNAFSTLPSLIKLDLSSNLLSSFPITGLHGLTHLKTGNHALQSLISSENFPPELKVIEMPTYAYQCC
	LGR6	QALDLSWNAIRSIHPEAFSTLHSLVKLDLTDNQLTTPLAGLGLMHKLKGNLALSQAFSKDSFPKLRILEVPYAYQCC
30	LGR4	RILDLSRNLIHEHSRAFATLGPITNLDVSFNELTSFPTEGLNGLNQLKIVGNFKLKEALAAKDFVNRLRSLSVPYAYQCC
	Consensus/80%	p.LsLtbN.I..IhspAfTL.s1hpLd1*.NbL**bPh.GL.GL.pLKL.GNb.Lpphbtc.FspL+.1phPYAYQCC
	LGR5	AFGVCEENAYKISNQWNKGDNSSMDD-LHKKDAGMFQAQDE---RDLEDFLLDFEEDLKALHSVQCSPSPGPFPKPCEHLL
	LGR6	PYGMCASFFKASGQWEAEDLHLDDEESSKRPLGLLARQAENHYDQDLDELQLEMEDS-KPHPSVQCSPTFPGPFPKPCEYLF
35	LGR4	AFWGCDSYANLN---TEDNSLQDHSAQEKGKCMSTADA---ANVTSTLENEEHQSIIHCTPSTGAFKPCEYLL
	Consensus/80%	sahhC.shhhphs.....D.p...Dc...pc.h.hb...s.....ph..pbEpp.+sb..1pC+p*sGsfKPCEaLb
	LGR5	DGWLIRIGVWTIAVLALTCNA-LVTSTVFR-SPLYISPIKLLIGVIAAVNMLTGVSSAVLAGVDAFTFGSEARHGAWWENG
	LGR6	ESWGIRLAVWAIVLISVLNCNGLVLTTVFAAGPVPLPPVKFVVGAIAGANTLTG1SCGLLASVDALTFGQFSEYGRWETG
40	LGR4	GSMWIRLTVWFIFLVALFFNLVLVLTTFASCTSLPSSKLFIGLISVSNLFMGIYTG1LTFLDAVSWGREFEFGIWWETG
	Consensus/80%	.twhIR1sVWhIh1lt1hhNhLVh.TsF..tsh.iss.Kbh1GhItssNhbhGl.st1Ls.1DAh*aGpFtcaGhbWEsG

continued as Fig. 4 (2/2)

Fig. 4 (2/2)

continued from Fig. 4 (1/2)

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LGR5	VGCHVIGFLSIFASESSVFLLTLAALERGFSVKYSAKFETKAPFSSLKVIIILCALLALTMAAVPLLGGSKYGA\$PLCLP
LGR6	LGCRATGFLAVLGSSEASVLLLTAAVQCSVSVCSRAGVKSPSLGSVRAGVLGCLALAGLAALPLASVGEYGASPLCLP
LGR4	SGCKVAGFLAVFSSESAIFLMLATVERSLSAKDIMKNGKSNHLKQFRVAALLAFLGATVAGCFPLFHRGEYSASPLCLP
Consensus/80%	.GC+shGFLtlbtSett1bLLhLa\$lpptbSsp.....pps.b.ph+shh1hshhhAhhtshPLh..tcYtASPLCLP

10

LGR5	--LPFGEPS TMGYMVALILLNSLCFLMMTIAYTKLYCNLDKGDL ENIWDCSMVKHIALLLFTNCILNCPVAFLSFSSLIN
LGR6	YAPPEGQPAALGFTVALVMMNSFCFLVVAGAYIKLYCDLPRGDFEAVWDCAMVRHVANLIFADGLLYCPCVAFLSFASMLG
LGR4	--FPTGETPSLGFVTIVLNLNSLAFLLMATVYT KLYCNLEKEDELSSENSQSSMIKHVAWLIFTNCIFFCPVAFSFAPLIT
Consensus/80%	...P.GpsssbGahVsL1bbNSbsFLhhshhYhKLYcsL.+Dbp...pstM1+H1AbL1Fsss1b.CPVAfbSfstsbls

15

LGR5	LTFISPEVIKFILLVVVPLPACLNPLLY1LFNPHEKEDLVSLRKQTYVWTR8KHPMSLMSINSDDVEKQSCDS-----TQ
LGR6	LFPVTPEAVKSVLLVLPACLNPLLYLLFNPHFRDDLRLRPR-----AGDSGPLAYAAAGELEKSSCD-----TQ
LGR4	AISISPEIMKSVTLLIFFPLPACLNPEVLYVFFNPKFEDWKLLKRRVTKSG9VSVISQGGCLEQDFYYDCGMYSHLQG
Consensus/80%	hh.1*PEhhHK.1hLlhhpLPA CLNPPLLY1b5NP+F+-Db..L+p.....t.pss1...sts..pc..hDs.....p.

20

LGR5	ALVTTTSSSITYDLPPS-----SVPSPAYFVTECHLS-----
LGR6	ALVAFSDVDLILEASEAGRPPGLETYGFPSTLISCQPGAPRLEGSHCVEPEGNHFGNPQPSMDGELLRAEGSTPAGG
LGR4	NLTVCDCCESFLLTkpVsCKHLIKSHSCPALAVASCQRPEGYWSDCGTQSAHSDYADEEDSFVSDSSDQVQACGRACFYQ
Consensus/80%	sLsshsssp.hb.h..s.....Ss.pP.h...-ssp.s.....

25

LGR5	-----SVAFVPCL
LGR6	GLSGGGGFQPSGLAFASHV
LGR4	SRGFPLVRYAYNLPRVKD-
Consensus/80%s1sbs.p.

30